

## WEST Search History

DATE: Tuesday, January 10, 2006

**Hide?** **Set Name Query** **Hit Count**

*DB=PGPB,USPT; PLUR=YES; OP=OR*

<input type="checkbox"/>	L2	VAN adj GROENINGHEN	3
--------------------------	----	---------------------	---

<input type="checkbox"/>	L1	VAN GROENINGHEN	277930
--------------------------	----	-----------------	--------

END OF SEARCH HISTORY

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:ssptacmb1647

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 SEP 09 ACD predicted properties enhanced in REGISTRY/ZREGISTRY  
NEWS 4 OCT 03 MATHDI removed from STN  
NEWS 5 OCT 04 CA/Caplus-Canadian Intellectual Property Office (CIPO) added  
to core patent offices  
NEWS 6 OCT 13 New CAS Information Use Policies Effective October 17, 2005  
NEWS 7 OCT 17 STN(R) AnaVist(TM), Version 1.01, allows the export/download  
of Caplus documents for use in third-party analysis and  
visualization tools  
NEWS 8 OCT 27 Free KWIC format extended in full-text databases  
NEWS 9 OCT 27 DIOGENES content streamlined  
NEWS 10 OCT 27 EPFULL enhanced with additional content  
NEWS 11 NOV 14 CA/Caplus - Expanded coverage of German academic research  
NEWS 12 NOV 30 REGISTRY/ZREGISTRY on STN(R) enhanced with experimental  
spectral property data  
NEWS 13 DEC 05 CASREACT(R) - Over 10 million reactions available  
NEWS 14 DEC 14 2006 MeSH terms loaded in MEDLINE/LMEDLINE  
NEWS 15 DEC 14 2006 MeSH terms loaded for MEDLINE file segment of TOXCENTER  
NEWS 16 DEC 14 CA/Caplus to be enhanced with updated IPC codes  
NEWS 17 DEC 16 MARPATprev will be removed from STN on December 31, 2005  
NEWS 18 DEC 21 IPC search and display fields enhanced in CA/Caplus with the  
IPC reform  
NEWS 19 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/USPAT2  
  
NEWS EXPRESS JANUARY 03 CURRENT VERSION FOR WINDOWS IS V8.01,  
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.  
V8.0 USERS CAN OBTAIN THE UPGRADE TO V8.01 AT  
<http://download.cas.org/express/v8.0-Discover/>  
  
NEWS DCOST SINCE APPROXIMATELY 20:00 COLUMBUS TIME DECEMBER 29,  
SOME ONLINE COST DISPLAYS HAVE BEEN SHOWING COSTS IN  
2006 PRICES FOR STN COLUMBUS FILES. THIS HAS BEEN  
CORRECTED. PLEASE BE ASSURED THAT YOU WILL BE BILLED  
ACCORDING TO 2005 PRICES UNTIL JAN 1. PLEASE CONTACT  
YOUR LOCAL HELP DESK IF YOU HAVE ANY QUESTIONS. WE  
APOLOGIZE FOR THE ERROR.  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that  
specific topic.

All use of STN is subject to the provisions of the STN Customer

agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

\* \* \* \* \* STN Columbus \* \* \* \* \*

\*IFICDB - The IFI Comprehensive Database from 1950-present  
\*IFIPAT - The IFI Patent Database from 1950-present  
\*IFIUDB - The IFI Uniterm Database from 1950-present

\* The files listed above are temporarily unavailable.

FILE 'HOME' ENTERED AT 09:19:28 ON 10 JAN 2006

=> file medline	SINCE FILE	TOTAL
COST IN U.S. DOLLARS	ENTRY	SESSION
FULL ESTIMATED COST	0.63	0.63

FILE 'MEDLINE' ENTERED AT 09:21:17 ON 10 JAN 2006

FILE LAST UPDATED: 7 JAN 2006 (20060107/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 will soon be available. For details on the 2005 reload, enter HELP RLOAD at an arrow prompt (=>).  
See also:

<http://www.nlm.nih.gov/mesh/>  
[http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\\_mesh.html](http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_med\\_data\\_changes.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_2006\\_MeSH.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html)

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate

```
=> s tu GnRH or gonadotropin(w)releasing(w)hormone
    1145754 TU
      204 TUS
    1145918 TU
          (TU OR TUS)
    11745 GNRH
      107 GNRHS
    11747 GNRH
          (GNRH OR GNRHS)
      0 TU GNRH
          (TU(W) GNRH)
    41701 GONADOTROPIN
    22719 GONADOTROPINS
    55507 GONADOTROPIN
          (GONADOTROPIN OR GONADOTROPINS)
    57465 RELEASING
    268964 HORMONE
    177879 HORMONES
    389158 HORMONE
          (HORMONE OR HORMONES)
      7655 GONADOTROPIN(W) RELEASING(W) HORMONE
L1      7655 TU GNRH OR GONADOTROPIN(W) RELEASING(W) HORMONE
```

```
=> s l1 and Kaposi(w)sarcoma
      10558 KAPOS I
        40 KAPOSIS
      10568 KAPOS I
          (KAPOS I OR KAPOSIS)
      70545 SARCOMA
      11081 SARCOMAS
        119 SARCOMATA
      73808 SARCOMA
          (SARCOMA OR SARCOMAS OR SARCOMATA)
      1041 KAPOS I(W) SARCOMA
L2      0 L1 AND KAPOS I(W) SARCOMA
```

```
=> s l1 and glioblastoma
      11148 GLIOBLASTOMA
      2325 GLIOBLASTOMAS
      12012 GLIOBLASTOMA
          (GLIOBLASTOMA OR GLIOBLASTOMAS)
L3      0 L1 AND GLIOBLASTOMA
```

```
=> s l1 and medulloblastoma
      4566 MEDULLOBLASTOMA
      1170 MEDULLOBLASTOMAS
      4907 MEDULLOBLASTOMA
          (MEDULLOBLASTOMA OR MEDULLOBLASTOMAS)
L4      6 L1 AND MEDULLOBLASTOMA
```

```
=> dup rem
ENTER L# LIST OR (END):14
PROCESSING COMPLETED FOR L4
L5      6 DUP REM L4 (0 DUPLICATES REMOVED)
```

```
=> dis ibib abs l5 1-6
```

```
L5  ANSWER 1 OF 6      MEDLINE on STN
ACCESSION NUMBER:      2005069367      MEDLINE
DOCUMENT NUMBER:      PubMed ID: 15562029
TITLE:      Differential role of progesterone receptor isoforms in the
transcriptional regulation of human gonadotropin-
releasing hormone I (GnRH I) receptor,
GnRH I, and GnRH II.
AUTHOR:      An Beum-Soo; Choi Jung-Hye; Choi Kyung-Chul; Leung Peter C
K
CORPORATE SOURCE:      Department of Obstetrics and Gynecology, University of
British Columbia, Vancouver, British Columbia, Canada V6H
3V5.
SOURCE:      Journal of clinical endocrinology and metabolism, (2005
Feb) 90 (2) 1106-13. Electronic Publication: 2004-11-23.
Journal code: 0375362. ISSN: 0021-972X.
PUB. COUNTRY:      United States
DOCUMENT TYPE:      Journal; Article; (JOURNAL ARTICLE)
LANGUAGE:      English
FILE SEGMENT:      Abridged Index Medicus Journals; Priority Journals
ENTRY MONTH:      200503
ENTRY DATE:      Entered STN: 20050209
Last Updated on STN: 20050325
Entered Medline: 20050324
AB  Hypothalamic GnRH is a decapeptide that plays a pivotal role in mammalian
reproduction by stimulating the synthesis and secretion of gonadotropins
via binding to the GnRH receptor on the pituitary gonadotropins. It is
hypothesized that sex steroids may regulate GnRH I (a classical form of
GnRH), GnRH II (a second form of GnRH), and GnRH I receptor (GnRHRI) at
the transcriptional level in target tissues. Thus, in the present study a
role for progesterone (P4) in the regulation of GnRH I, GnRH II, and
```

GnRHRI was investigated using a human neuronal medulloblastoma cell line (TE671) as an in vitro model. The cells were transfected with human GnRHRI promoter-luciferase constructs, and promoter activities were analyzed after P4 treatment by luciferase and beta-galactosidase assay. The mRNA levels of GnRH I and GnRH II were analyzed by RT-PCR. Treatment of TE671 cells with P4 resulted in a decrease in GnRHRI promoter activity compared with the control level in a dose- and time-dependent manner. Cotreatment of these cells with RU486, an antagonist of P4, reversed P4-induced inhibition of GnRHRI promoter activity, suggesting that the P4 effect is mediated by P4 receptor (PR). In the cells transfected with a full-length of PR A- or PR B-expressing vector, overexpression of PR A increased the sensitivity toward P4 in an inhibition of GnRHRI promoter, whereas PR B increased transcriptional activity of GnRHRI promoter in the presence of P4. However, PR B itself did not act as a transcriptional activator of GnRHRI promoter. Because TE671 cells have been recently demonstrated to express and synthesize two forms of GnRHs, we also investigated the regulation of GnRH mRNAs by P4. In the present study, P4 increased GnRH I mRNA levels in a time- and dose-dependent manner. This stimulatory effect of P4 in the regulation of GnRH I mRNAs was significantly attenuated by RU486, whereas no significant difference in the expression level of GnRH II was observed with P4 or RU486. Interestingly, although the expression level of PR B was low compared with that of PR A, P4 action on the GnRH I gene was mediated by PR B. In conclusion, these results indicate that P4 is a potent regulator of GnRHRI at the transcriptional level as well as GnRH I mRNA. This distinct effect of P4 on the GnRH system may be derived from different pathways through PR A or PR B.

L5 ANSWER 2 OF 6 MEDLINE on STN  
 ACCESSION NUMBER: 2002198301 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 11931351  
 TITLE: The transcription of the hGnRH-I and hGnRH-II genes in human neuronal cells is differentially regulated by estrogen.  
 AUTHOR: Chen Alon; Zi Keren; Laskar-Levy Orly; Koch Yitzhak  
 CORPORATE SOURCE: Department of Neurobiology, Weizmann Institute of Science, Rehovot, Israel.  
 SOURCE: Journal of molecular neuroscience : MN, (2002 Feb-Apr) 18 (1-2) 67-76.  
 Journal code: 9002991. ISSN: 0895-8696.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 200211  
 ENTRY DATE: Entered STN: 20020405  
 Last Updated on STN: 20021211  
 Entered Medline: 20021108

AB Gonadotropin releasing hormone-I (GnRH-I), a decapeptide serves as a key regulator of reproduction. Recently, several groups have identified in the mammalian brain a second form of GnRH, of unknown function, designated GnRH-II. The human neuronal medulloblastoma cells (TE-671) were recently demonstrated to express the two forms of GnRH (GnRH-I and GnRH-II). We used this cell line, as a model system, to investigate the regulation of human GnRH-I and GnRH-II genes by estrogen. Estrogen is one of the principal regulators of GnRH-I in hypothalamic neurons, acting as a classic homeostatic feedback molecule between the gonads and the brain. In this study, we investigated the regulation of the two GnRH forms by estrogen, in the human neuronal cell line TE-671. We demonstrate, for the first time, that the hGnRH-II and hGnRH-I genes are differentially regulated by estrogen. Using reverse transcriptase-polymerase chain reaction (RT-PCR) and Southern hybridization, we found that estrogen increases endogenous hGnRH-II mRNA levels and decreases endogenous hGnRH-I mRNA levels. Furthermore, we found these effects to be promoter-mediated. We cloned the hGnRH-I and

hGnRH-II promoter constructs upstream to a luciferase reporter plasmid, and cotransfected these constructs with an estrogen receptor alpha into the TE-671 neuronal cells. Luciferase activity of GnRH promoter constructs treated with estrogen demonstrates that the differential regulation of the GnRH genes by estrogen is mediated at the transcription level.

L5 ANSWER 3 OF 6 MEDLINE on STN  
 ACCESSION NUMBER: 2001179519 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 11159856  
 TITLE: Two isoforms of gonadotropin-releasing hormone are coexpressed in neuronal cell lines.  
 AUTHOR: Chen A; Yahalom D; Laskar-Levy O; Rahimipour S; Ben-Aroya N; Koch Y  
 CORPORATE SOURCE: Department of Neurobiology, Weizmann Institute of Science, Rehovot 76100, Israel.  
 SOURCE: Endocrinology, (2001 Feb) 142 (2) 830-7.  
 Journal code: 0375040. ISSN: 0013-7227.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals  
 ENTRY MONTH: 200103  
 ENTRY DATE: Entered STN: 20010404  
 Last Updated on STN: 20010404  
 Entered Medline: 20010329

AB GnRH-I serves as the neuropeptide that regulates mammalian reproduction. Recently, several groups have identified in the brain of rodents, monkeys, and humans a second isoform of GnRH (GnRH-II) whose structure is 70% identical to that of GnRH-I. In this study we demonstrate for the first time human and mouse neuronal cell lines that express both GnRH-I and GnRH-II. Following the screening of several human neuronal cell lines by RT-PCR and Southern hybridization, we demonstrated that two cell lines, TE-671 medulloblastoma and LAN-1 neuroblastoma cells, coexpress messenger RNA encoding the two isoforms of GnRH. Nucleotide sequencing indicated that the complementary DNA fragments are identical to those of the known human GnRH-I and GnRH-II sequences. Extracts obtained from the TE-671 and LAN-1 cell lines as well as from the immortalized mouse hypothalamic GT1-7 neuronal cell line were found to contain the two isoforms of GnRH, which exhibited identical chromatographic properties as synthetic GnRH-I and GnRH-II, in HPLC followed by specific RIAs. Furthermore, double immunofluorescence studies demonstrated the two GnRH isoforms in LAN-1, TE-671, and GT1-7 cells. The identification of neuronal cell lines expressing both GnRH-I and GnRH-II provides tools for studying the differential regulation of gene expression and secretion and for studying the interaction between the two isoforms. Such studies may contribute to elucidation of the physiological functions of GnRH-II, which are still unknown.

L5 ANSWER 4 OF 6 MEDLINE on STN  
 ACCESSION NUMBER: 2001050966 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 11033292  
 TITLE: A case of atypical absence seizures induced by leuprolide acetate.  
 AUTHOR: Akaboshi S; Takeshita K  
 CORPORATE SOURCE: Division of Child Neurology, Institute of Neurological Sciences, Faculty of Medicine, Tottori University, Yonago, Japan.  
 SOURCE: Pediatric neurology, (2000 Sep) 23 (3) 266-8.  
 Journal code: 8508183. ISSN: 0887-8994.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: (CASE REPORTS)  
 Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals

ENTRY MONTH: 200012  
ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20001211

AB We report a case of a 13-year-old female with atypical absence seizures induced by prolonged administration of long-acting leuprolide acetate (LA). This patient had brain involvement resulting from chemotherapy and radiotherapy for a medulloblastoma. At 13 years of age, administration of long-acting LA was started. After the third dose of long-acting LA, atypical absence seizures appeared. After discontinuing long-acting LA, the seizures stopped without administration of any antiepileptic drugs. However, 2 years, 6 months later, the same seizures again appeared. On the basis of the findings of endocrinologic investigations and the reported data of pharmacokinetics of LA, we speculate that her seizures were induced by LA and that the seizures were associated with the presence of brain damage in the patient. Care should therefore be taken when using long-acting LA or other gonadotropin-releasing hormone analogues for pediatric patients with diffuse brain damage.

L5 ANSWER 5 OF 6 MEDLINE on STN  
ACCESSION NUMBER: 2000079233 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 10611579  
TITLE: Adult height after growth hormone (GH) treatment for GH deficiency due to cranial irradiation.  
AUTHOR: Adan L; Sainte-Rose C; Souberbielle J C; Zucker J M; Kalifa C; Brauner R  
CORPORATE SOURCE: Pediatric Endocrinology Department, Universite Rene Descartes and Hopital Necker-Enfants Malades, Assistance Publique-Hopitaux de Paris, Paris, France.  
SOURCE: Medical and pediatric oncology, (2000 Jan) 34 (1) 14-9.  
Journal code: 7506654. ISSN: 0098-1532.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: (CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200002  
ENTRY DATE: Entered STN: 20000209  
Last Updated on STN: 20000209  
Entered Medline: 20000203

AB BACKGROUND: The indications and factors affecting the growth in response to treatment with growth hormone (GH) of patients with cranial irradiation-induced GH deficiency remain unclear. PROCEDURE: The adult heights of 56 patients treated with GH (0.4-0.6 U/kg/week) as daily sc injections were analysed. They had been given 18 or 24 Grays (Gy) cranial irradiation for leukemia (group 1, 26 cases), 50 +/- 1 Gy for various tumors (group 2, 13 cases), 46 +/- 1 Gy for retinoblastoma (group 3, 8 cases), or 34 +/- 2 Gy with spinal irradiation for medulloblastoma (group 4, 9 cases). Twenty-five of these 56 patients had early puberty and were also treated with gonadotropin-releasing hormone (GnRH) analog. RESULTS: The standing (-1.0 +/- 0.2 in group 1, -0.7 +/- 0.3 in group 2, -1.1 +/- 0.3 in group 3, and -2.0 +/- 0.4 SD in group 4) and sitting (-1.8 +/- 0.2 in group 1, -0.4 +/- 0.4 in group 2, -1.2 +/- 0.4 in group 3, and -3.4 +/- 0.4 SD in group 4) adult heights were shorter (P < 0.05 for standing and P < 0.001 for sitting heights) for group 4 than for each of the other groups. Of the 47 patients given cranial (and not craniospinal) irradiation, sitting adult height was shorter (P = 0.02) and the difference between standing adult and target heights greater (P = 0.03) in those patients in whom puberty occurred at a normal age than in those treated with GnRH analog. Conclusion. The incomplete catch-up of growth seems to be mainly due to the reduction in sitting height of patients given spinal irradiation and in whom puberty occurred at a normal age. This suggests that GnRH analog treatment should be more widely used to treat children with early and/or

rapidly progressing puberty after cranial irradiation.  
Copyright 2000 Wiley-Liss, Inc.

L5 ANSWER 6 OF 6 MEDLINE on STN  
ACCESSION NUMBER: 1999258039 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 10326189  
TITLE: Effects of puberty on bone age maturation in a girl after  
medulloblastoma therapy.  
AUTHOR: Marx M; Schoof E; Grabenbauer G G; Beck J D; Doerr H G  
CORPORATE SOURCE: Division of Paediatric Endocrinology, University of  
Erlangen-Nuremberg, Germany.  
SOURCE: Journal of pediatric and adolescent gynecology, (1999 May)  
12 (2) 62-6.  
Journal code: 9610774. ISSN: 1083-3188.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: (CASE REPORTS)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199906  
ENTRY DATE: Entered STN: 19990714  
Last Updated on STN: 19990714  
Entered Medline: 19990629

AB BACKGROUND: Craniospinal radiotherapy for malignant brain tumors can  
result in a variety of neuroendocrine disturbances, among which are the  
development of growth hormone deficiency and early puberty, which can  
markedly reduce adult height. METHODS: The authors report the case of a  
girl who received craniospinal radiotherapy for a medulloblastoma  
at the age of 3.4 years. At 9.1 years, growth hormone therapy was  
started, and spontaneous onset of puberty (Tanner stage B2) occurred at  
age 10.3 years. Interval until menarche was short, at only 0.9 years.  
RESULTS: Although chronologic age at appearance of Tanner stages was  
within the normal range, the patient showed a rapid acceleration in  
skeletal maturation, resulting in adult short stature. CONCLUSION: Bone  
age seems to be a more precise parameter for biologic maturation in some  
patients after craniospinal irradiation than is clinical assessment of  
pubertal stages. Thus, if progression of bone age and decreasing final  
height predictions are noted, puberty should be stopped with  
gonadotropin-releasing hormone analogs, even  
if pubertal development seems to be adequate for chronologic age, because  
this increases the remaining time for growth hormone treatment.

=>

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	13.02	13.65

STN INTERNATIONAL LOGOFF AT 09:39:24 ON 10 JAN 2006

Connecting via Winsock to STN



Welcome to STN International! Enter x:x

LOGINID:ssptacmb1647

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 SEP 09 ACD predicted properties enhanced in REGISTRY/ZREGISTRY  
NEWS 4 OCT 03 MATHDI removed from STN  
NEWS 5 OCT 04 CA/CAPplus-Canadian Intellectual Property Office (CIPO) added  
to core patent offices  
NEWS 6 OCT 13 New CAS Information Use Policies Effective October 17, 2005  
NEWS 7 OCT 17 STN(R) AnaVist(TM), Version 1.01, allows the export/download  
of CAPplus documents for use in third-party analysis and  
visualization tools  
NEWS 8 OCT 27 Free KWIC format extended in full-text databases  
NEWS 9 OCT 27 DIOGENES content streamlined  
NEWS 10 OCT 27 EPFULL enhanced with additional content  
NEWS 11 NOV 14 CA/CAPplus - Expanded coverage of German academic research  
NEWS 12 NOV 30 REGISTRY/ZREGISTRY on STN(R) enhanced with experimental  
spectral property data  
NEWS 13 DEC 05 CASREACT(R) - Over 10 million reactions available  
NEWS 14 DEC 14 2006 MeSH terms loaded in MEDLINE/LMEDLINE  
NEWS 15 DEC 14 2006 MeSH terms loaded for MEDLINE file segment of TOXCENTER  
NEWS 16 DEC 14 CA/CAPplus to be enhanced with updated IPC codes  
NEWS 17 DEC 16 MARPATprev will be removed from STN on December 31, 2005  
NEWS 18 DEC 21 IPC search and display fields enhanced in CA/CAPplus with the  
IPC reform  
NEWS 19 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/USPAT2  
  
NEWS EXPRESS JANUARY 03 CURRENT VERSION FOR WINDOWS IS V8.01,  
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.  
V8.0 USERS CAN OBTAIN THE UPGRADE TO V8.01 AT  
<http://download.cas.org/express/v8.0-Discover/>  
  
NEWS DCOST SINCE APPROXIMATELY 20:00 COLUMBUS TIME DECEMBER 29,  
SOME ONLINE COST DISPLAYS HAVE BEEN SHOWING COSTS IN  
2006 PRICES FOR STN COLUMBUS FILES. THIS HAS BEEN  
CORRECTED. PLEASE BE ASSURED THAT YOU WILL BE BILLED  
ACCORDING TO 2005 PRICES UNTIL JAN 1. PLEASE CONTACT  
YOUR LOCAL HELP DESK IF YOU HAVE ANY QUESTIONS. WE  
APOLOGIZE FOR THE ERROR.  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that  
specific topic.

All use of STN is subject to the provisions of the STN Customer  
agreement. Please note that this agreement limits use to scientific  
research. Use for software development or design or implementation  
of commercial gateways or other similar uses is prohibited and may

result in loss of user privileges and other penalties.

\*\*\*\*\* STN Columbus \*\*\*\*\*

\*IFICDB - The IFI Comprehensive Database from 1950-present  
\*IFIPAT - The IFI Patent Database from 1950-present  
\*IFIUDB - The IFI Uniterm Database from 1950-present

\* The files listed above are temporarily unavailable.

FILE 'HOME' ENTERED AT 09:40:00 ON 10 JAN 2006

=> file medline

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'MEDLINE' ENTERED AT 09:40:08 ON 10 JAN 2006

FILE LAST UPDATED: 7 JAN 2006 (20060107/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 will soon be available. For details on the 2005 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

<http://www.nlm.nih.gov/mesh/>  
[http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\\_mesh.html](http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_med\\_data\\_changes.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_2006\\_MeSH.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html)

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate

=> s l1 and pinealoma

L1 NOT FOUND

The L-number entered could not be found. To see the definition of L-numbers, enter DISPLAY HISTORY at an arrow prompt (=>).

=> logoff

ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.39	0.60

STN INTERNATIONAL LOGOFF AT 09:40:46 ON 10 JAN 2006

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:ssptacmb1647

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 SEP 09 ACD predicted properties enhanced in REGISTRY/ZREGISTRY  
NEWS 4 OCT 03 MATHDI removed from STN  
NEWS 5 OCT 04 CA/Caplus-Canadian Intellectual Property Office (CIPO) added  
to core patent offices  
NEWS 6 OCT 13 New CAS Information Use Policies Effective October 17, 2005  
NEWS 7 OCT 17 STN(R) AnaVist(TM), Version 1.01, allows the export/download  
of Caplus documents for use in third-party analysis and  
visualization tools  
NEWS 8 OCT 27 Free KWIC format extended in full-text databases  
NEWS 9 OCT 27 DIOGENES content streamlined  
NEWS 10 OCT 27 EPFULL enhanced with additional content  
NEWS 11 NOV 14 CA/Caplus - Expanded coverage of German academic research  
NEWS 12 NOV 30 REGISTRY/ZREGISTRY on STN(R) enhanced with experimental  
spectral property data  
NEWS 13 DEC 05 CASREACT(R) - Over 10 million reactions available  
NEWS 14 DEC 14 2006 MeSH terms loaded in MEDLINE/LMEDLINE  
NEWS 15 DEC 14 2006 MeSH terms loaded for MEDLINE file segment of TOXCENTER  
NEWS 16 DEC 14 CA/Caplus to be enhanced with updated IPC codes  
NEWS 17 DEC 16 MARPATprev will be removed from STN on December 31, 2005  
NEWS 18 DEC 21 IPC search and display fields enhanced in CA/Caplus with the  
IPC reform  
NEWS 19 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/USPAT2  
  
NEWS EXPRESS JANUARY 03 CURRENT VERSION FOR WINDOWS IS V8.01,  
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.  
V8.0 USERS CAN OBTAIN THE UPGRADE TO V8.01 AT  
<http://download.cas.org/express/v8.0-Discover/>  
  
NEWS DCOST SINCE APPROXIMATELY 20:00 COLUMBUS TIME DECEMBER 29,  
SOME ONLINE COST DISPLAYS HAVE BEEN SHOWING COSTS IN  
2006 PRICES FOR STN COLUMBUS FILES. THIS HAS BEEN  
CORRECTED. PLEASE BE ASSURED THAT YOU WILL BE BILLED  
ACCORDING TO 2005 PRICES UNTIL JAN 1. PLEASE CONTACT  
YOUR LOCAL HELP DESK IF YOU HAVE ANY QUESTIONS. WE  
APOLOGIZE FOR THE ERROR.  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that  
specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

\* \* \* \* \* STN Columbus \* \* \* \* \*

\*IFICDB - The IFI Comprehensive Database from 1950-present  
\*IFIPAT - The IFI Patent Database from 1950-present  
\*IFIUDB - The IFI Uniterm Database from 1950-present

\* The files listed above are temporarily unavailable.

FILE 'HOME' ENTERED AT 09:42:34 ON 10 JAN 2006

=> file medline	SINCE FILE	TOTAL
COST IN U.S. DOLLARS	ENTRY	SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'MEDLINE' ENTERED AT 09:42:59 ON 10 JAN 2006

FILE LAST UPDATED: 7 JAN 2006 (20060107/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 will soon be available. For details on the 2005 reload, enter HELP RLOAD at an arrow prompt (=>).  
See also:

<http://www.nlm.nih.gov/mesh/>  
[http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\\_mesh.html](http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_med\\_data\\_changes.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_2006\\_MeSH.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html)

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate

=> s tu GnRH or gonadotropin(w)releasing(w)hormone  
1145754 TU  
204 TUS  
1145918 TU  
(TU OR TUS)  
11745 GNRH  
107 GNRHS  
11747 GNRH  
(GNRH OR GNRHS)  
0 TU GNRH  
(TU(W)GNRH)  
41701 GONADOTROPIN  
22719 GONADOTROPINS  
55507 GONADOTROPIN  
(GONADOTROPIN OR GONADOTROPINS)  
57465 RELEASING  
268964 HORMONE  
177879 HORMONES  
389158 HORMONE  
(HORMONE OR HORMONES)

```

7655 GONADOTROPIN(W) RELEASING(W) HORMONE
L1      7655 TU GNRH OR GONADOTROPIN(W) RELEASING(W) HORMONE

=> s l1 and pinealoma
      1238 PINEALOMA
      51 PINEALOMAS
      1251 PINEALOMA
          (PINEALOMA OR PINEALOMAS)
L2      0 L1 AND PINEALOMA

=> s l1 and neuroblastoma
      22738 NEUROBLASTOMA
      2006 NEUROBLASTOMAS
      23167 NEUROBLASTOMA
          (NEUROBLASTOMA OR NEUROBLASTOMAS)
L3      3 L1 AND NEUROBLASTOMA

=> dis ibib abs l2 1-3
L2 HAS NO ANSWERS
L1      7655 SEA FILE=MEDLINE ABB=ON PLU=ON TU GNRH OR GONADOTROPIN(W) RELE
      ASING(W) HORMONE
L2      0 SEA FILE=MEDLINE ABB=ON PLU=ON L1 AND PINEALOMA

=> dis ibib abs l3 1-3

L3 ANSWER 1 OF 3 MEDLINE on STN
ACCESSION NUMBER: 2003030963 MEDLINE
DOCUMENT NUMBER: PubMed ID: 12538601
TITLE: A transcriptionally active human type II
      gonadotropin-releasing hormone
      receptor gene homolog overlaps two genes in the antisense
      orientation on chromosome 1q.12.
AUTHOR: Morgan Kevin; Conklin Darrell; Pawson Adam J; Sellar Robin;
      Ott Thomas R; Millar Robert P
CORPORATE SOURCE: Medical Research Council Human Reproductive Sciences Unit,
      University of Edinburgh Academic Centre, Edinburgh EH16
      4SB, United Kingdom.. k.morgan@hrrsu.mrc.ac.uk
SOURCE: Endocrinology, (2003 Feb) 144 (2) 423-36.
      Journal code: 0375040. ISSN: 0013-7227.
PUB. COUNTRY: United States
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: English
FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals
ENTRY MONTH: 200302
ENTRY DATE: Entered STN: 20030123
      Last Updated on STN: 20030214
      Entered Medline: 20030213
AB GnRH-II peptide hormone exhibits complete sequence conservation across
vertebrate species, including man. Type-II GnRH receptor genes have been
characterized recently in nonhuman primates, but the human receptor gene
homolog contains a frameshift, a premature stop codon (UGA), and a 3'
overlap of the RBM8A gene on chromosome 1q.12. A retrotransposed
pseudogene, RBM8B, retains partial receptor sequence. In this study,
bioinformatics show that the human receptor gene promoter overlaps the
peroxisomal protein 11-beta gene promoter and the premature UGA is
positionally conserved in chimpanzee. A CGA [arginine (Arg)] occurs in
porcine DNA, but UGA is shifted one codon to the 5' direction in bovine
DNA, suggesting independent evolution of premature stop codons. In
contrast to marmoset tissue RNA, exon- and strand-specific probes are
required to distinguish differently spliced human receptor gene
transcripts in cell lines (HP75, IMR-32). RBM8B is not transcribed.
Sequencing of cDNAs for spliced receptor mRNAs showed no evidence for
alteration of the premature UGA by RNA editing, but alternative splicing

```

circumvents the frameshift to encode a two-membrane-domain protein before this UGA. A stem-loop motif resembling a selenocysteine insertion sequence and a potential alternative translation initiation site might enable expression of further proteins involved in interactions within the GnRH system.

L3 ANSWER 2 OF 3 MEDLINE on STN  
ACCESSION NUMBER: 2001179519 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 11159856  
TITLE: Two isoforms of gonadotropin-releasing hormone are coexpressed in neuronal cell lines.  
AUTHOR: Chen A; Yahalom D; Laskar-Levy O; Rahimipour S; Ben-Aroya N; Koch Y  
CORPORATE SOURCE: Department of Neurobiology, Weizmann Institute of Science, Rehovot 76100, Israel.  
SOURCE: Endocrinology, (2001 Feb) 142 (2) 830-7.  
Journal code: 0375040. ISSN: 0013-7227.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals  
ENTRY MONTH: 200103  
ENTRY DATE: Entered STN: 20010404  
Last Updated on STN: 20010404  
Entered Medline: 20010329

AB GnRH-I serves as the neuropeptide that regulates mammalian reproduction. Recently, several groups have identified in the brain of rodents, monkeys, and humans a second isoform of GnRH (GnRH-II) whose structure is 70% identical to that of GnRH-I. In this study we demonstrate for the first time human and mouse neuronal cell lines that express both GnRH-I and GnRH-II. Following the screening of several human neuronal cell lines by RT-PCR and Southern hybridization, we demonstrated that two cell lines, TE-671 medulloblastoma and LAN-1 neuroblastoma cells, coexpress messenger RNA encoding the two isoforms of GnRH. Nucleotide sequencing indicated that the complementary DNA fragments are identical to those of the known human GnRH-I and GnRH-II sequences. Extracts obtained from the TE-671 and LAN-1 cell lines as well as from the immortalized mouse hypothalamic GT1-7 neuronal cell line were found to contain the two isoforms of GnRH, which exhibited identical chromatographic properties as synthetic GnRH-I and GnRH-II, in HPLC followed by specific RIAs. Furthermore, double immunofluorescence studies demonstrated the two GnRH isoforms in LAN-1, TE-671, and GT1-7 cells. The identification of neuronal cell lines expressing both GnRH-I and GnRH-II provides tools for studying the differential regulation of gene expression and secretion and for studying the interaction between the two isoforms. Such studies may contribute to elucidation of the physiological functions of GnRH-II, which are still unknown.

L3 ANSWER 3 OF 3 MEDLINE on STN  
ACCESSION NUMBER: 81142294 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 6259147  
TITLE: Interaction of fluorescent gonadotropin-releasing hormone with receptors in cultured pituitary cells.  
AUTHOR: Naor Z; Atlas D; Clayton R N; Forman D S; Amsterdam A; Catt K J  
SOURCE: Journal of biological chemistry, (1981 Mar 25) 256 (6) 3049-52.  
Journal code: 2985121R. ISSN: 0021-9258.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 198105

ENTRY DATE: Entered STN: 19900316  
Last Updated on STN: 19970203  
Entered Medline: 19810526

AB A fluorescent derivative of the gonadotropin-releasing hormone (GnRH) agonist analog, [D-Lys6]GnRH, was synthesized for receptor studies and shown to be biologically active. The rhodamine-derivatized peptide (Rh-GnRH) retained 40% of the receptor binding activity of [D-Lys6]GnRH, and 50% of the luteinizing hormone-releasing activity assayed in cultured pituitary cells. The fluorescent analog was employed to visualize the distribution of GnRH receptors in cultured pituitary cells, using the technique of video-intensified fluorescence microscopy. The binding of Rh-GnRH was confined to the large gonadotrophs which comprised 15% of the cell population. The specificity of the binding was shown by the absence of significant fluorescence in the presence of a 100-fold excess of [D-Lys6]GnRH, or when Rh-GnRH was incubated with choriocarcinoma, neuroblastoma, or 3T3 cell lines devoid of GnRH receptors. The interaction of Rh-GnRH with living pituitary cells was characterized by an initial diffuse distribution, followed by the formation of polar aggregates that later appeared to be internalized. These observations emphasize the value of fluorescent derivatives of GnRH for elucidating the course of the interaction with specific receptors on pituitary gonadotrophs. The initial results indicate that GnRH-receptor complexes undergo aggregation during stimulation of luteinizing hormone release, and are later internalized for subsequent degradation and/ or intracellular actions.

=> s ll and craniopharyngeoma  
24 CRANIOPHARYNGEOMA  
20 CRANIOPHARYNGEOMAS  
43 CRANIOPHARYNGEOMA  
(CRANIOPHARYNGEOMA OR CRANIOPHARYNGEOMAS)  
L4 1 L1 AND CRANIOPHARYNGEOMA

=> dis ibib abs l4

L4 ANSWER 1 OF 1 MEDLINE on STN  
ACCESSION NUMBER: 91141849 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 1996204  
TITLE: [Pulsatile gonadotropin-releasing hormone substitution following excision of a craniopharyngioma with suprasellar invasion]. Pulzatorikus gonadotropin releasing hormon substitutio suprasellaris novekedesu craniopharyngeoma eltavolitasat kovetoen.  
AUTHOR: Koloszar S; Bartfai G; Sas M  
CORPORATE SOURCE: Szuleszeti es Nagygyaszati Klinika, Szent-Gyorgi Albert Orvostudományi Egyetem, Szeged.  
SOURCE: Orvosi hetilap, (1991 Jan 20) 132 (3) 139-41. Journal code: 0376412. ISSN: 0030-6002.  
PUB. COUNTRY: Hungary  
DOCUMENT TYPE: (CASE REPORTS)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: Hungarian  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199103  
ENTRY DATE: Entered STN: 19910412  
Last Updated on STN: 19910412  
Entered Medline: 19910326

AB Craniopharyngeoma growing suprasellary attacks the medio-basal region of hypothalamus, that leads to the stopping of the production of gonadotropin releasing hormone. In connection with the case of a 15-year-old girl who had partial extirpation of

craniopharyngeoma the authors write about the favourable endocrine effect of pulsatile gonadotropin releasing hormone treatment. Through giving gonadotropin releasing hormone every 90 minutes in 20 micrograms doses menstruation cycle and ovulation was performed. Beside surgical treatment hormonal substitution plays an important role in the treatment of additional endocrine symptoms.

```
=> s l1 and meningeoma
    93 MENINGEOMA
    56 MENINGEOMAS
    141 MENINGEOMA
        (MENINGEOMA OR MENINGEOMAS)
L5      0 L1 AND MENINGEOMA

=> s l1 and chordoma
    2211 CHORDOMA
    636 CHORDOMAS
    2301 CHORDOMA
        (CHORDOMA OR CHORDOMAS)
L6      0 L1 AND CHORDOMA

=> s l1 and Ewing sarcoma
    5448 EWING
    54 EWINGS
    5483 EWING
        (EWING OR EWINGS)
    70545 SARCOMA
    11081 SARCOMAS
    119 SARCOMATA
    73808 SARCOMA
        (SARCOMA OR SARCOMAS OR SARCOMATA)
    769 EWING SARCOMA
        (EWING(W) SARCOMA)
L7      0 L1 AND EWING SARCOMA

=> s l1 and malignant(w)melanoma
    191227 MALIGNANT
    9 MALIGNANTS
    191228 MALIGNANT
        (MALIGNANT OR MALIGNANTS)
    60376 MELANOMA
    9262 MELANOMAS
    80 MELANOMATA
    1 MELANOMATAS
    61358 MELANOMA
        (MELANOMA OR MELANOMAS OR MELANOMATA OR MELANOMATAS)
    16936 MALIGNANT(W) MELANOMA
L8      0 L1 AND MALIGNANT(W) MELANOMA

=> s l1 and Kaposi(w)sarcoma
    10558 KAPOSI
    40 KAPOSIS
    10568 KAPOSI
        (KAPOSI OR KAPOSIS)
    70545 SARCOMA
    11081 SARCOMAS
    119 SARCOMATA
    73808 SARCOMA
        (SARCOMA OR SARCOMAS OR SARCOMATA)
    1041 KAPOSI(W) SARCOMA
L9      0 L1 AND KAPOSI(W) SARCOMA
```



```
=> s l1 and (brain(w)tumor or meningeal(w)tumor)
661971 BRAIN
26207 BRAINS
667265 BRAIN
      (BRAIN OR BRAINS)
625474 TUMOR
271064 TUMORS
747795 TUMOR
      (TUMOR OR TUMORS)
14457 BRAIN(W)TUMOR
18967 MENINGEAL
1      MENINGEALS
18967 MENINGEAL
      (MENINGEAL OR MENINGEALS)
625474 TUMOR
271064 TUMORS
747795 TUMOR
      (TUMOR OR TUMORS)
131 MENINGEAL(W)TUMOR
L10      7 L1 AND (BRAIN(W)TUMOR OR MENINGEAL(W)TUMOR)
```

```
=> dup rem
ENTER L# LIST OR (END):l10
PROCESSING COMPLETED FOR L10
L11      7 DUP REM L10 (0 DUPLICATES REMOVED)
```

```
=> dis ibib abs l11 1-7
```

```
L11 ANSWER 1 OF 7      MEDLINE on STN
ACCESSION NUMBER:      2002464597      MEDLINE
DOCUMENT NUMBER:      PubMed ID: 12182973
TITLE:      Preirradiation endocrinopathies in pediatric brain
      tumor patients determined by dynamic tests of
      endocrine function.
AUTHOR:      Merchant Thomas E; Williams Tani; Smith Julie M; Rose Susan
      R; Danish Robert K; Burghen George A; Kun Larry E; Lustig
      Robert H
CORPORATE SOURCE:      Department of Radiation Oncology, St. Jude Children's
      Research Hospital, 332 North Lauderdale Street, Memphis, TN
      38105, USA.. thomas.merchant@stjude.org
CONTRACT NUMBER:      P30 CA 21765 (NCI)
SOURCE:      International journal of radiation oncology, biology,
      physics, (2002 Sep 1) 54 (1) 45-50.
      Journal code: 7603616. ISSN: 0360-3016.
PUB. COUNTRY:      United States
DOCUMENT TYPE:      Journal; Article; (JOURNAL ARTICLE)
LANGUAGE:      English
FILE SEGMENT:      Priority Journals
ENTRY MONTH:      200209
ENTRY DATE:      Entered STN: 20020913
      Last Updated on STN: 20020927
      Entered Medline: 20020926
AB PURPOSE: To prospectively evaluate pediatric patients with localized
      primary brain tumors for evidence of endocrinopathy
      before radiotherapy (RT). METHODS AND MATERIALS: Seventy-five pediatric
      patients were evaluated with the arginine tolerance test and L-dopa test
      for growth hormone secretory capacity and activity; thyroid-stimulating
      hormone surge and thyrotropin-releasing hormone stimulation test for the
      hypothalamic-thyroid axis; the 1-microg adrenocorticotropin hormone (ACTH)
      and metyrapone test for ACTH reserve; and, depending on age, a
      gonadotropin-releasing hormone stimulation
      test to determine gonadotropin response. The study included 38 male and
      37 female patients, age 1-21 years with ependymoma (n = 35), World Health
      Organization (WHO) Grade I-II astrocytoma (n = 18), WHO Grade III-IV
```

astrocytoma (n = 10), craniopharyngioma (n = 7), optic pathway tumor (n = 4), and germinoma (n = 1). Seven patients receiving dexamethasone at the time of the evaluation were excluded from the final analysis. RESULTS: Of 68 assessable patient, 45 (66%) had evidence of endocrinopathy before RT, including 15 of 32 patients (47%) with posterior fossa tumors. Of the 45 patients, 38% had growth hormone deficiency, 43% had thyroid-stimulating hormone secretion abnormality, 22% had an abnormality in ACTH reserve, and 13% had an abnormality in age-dependent gonadotropin secretion. CONCLUSION: The incidence of pre-RT endocrinopathy in pediatric brain tumor patients is high, including patients with tumors not adjacent to the hypothalamic-pituitary unit. These data suggest an overestimation in the incidence of radiation-induced endocrinopathy. Baseline endocrine function should be determined for brain tumor patients before therapy. The potential for radiation-induced endocrinopathy alone cannot be used as an argument for alternatives to RT for most patients. Pre-RT endocrinopathy may be an early indicator of central nervous system damage that will influence the functional outcome unrelated to RT.

L11 ANSWER 2 OF 7 MEDLINE on STN  
 ACCESSION NUMBER: 2001518088 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 11528557  
 TITLE: [Cranial irradiation induces premature activation of the gonadotropin-releasing-hormone ].  
 Schadelbestrahlung verursacht vorzeitige Aktivierung des Gonadotropin-Releasinhormon (GnRH)-Pulsgenerators bei Ratten - Ein neues Tiermodell fur strahleninduzierte Storungen der Pubertat.  
 AUTHOR: Roth C; Lakomek M; Schmidberger H; Jarry H  
 CORPORATE SOURCE: Kinderklinik, Germany.  
 SOURCE: Klinische Padiatrie, (2001 Jul-Aug) 213 (4) 239-43.  
 Journal code: 0326144. ISSN: 0300-8630.  
 PUB. COUNTRY: Germany; Germany, Federal Republic of  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: German  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 200112  
 ENTRY DATE: Entered STN: 20010924  
 Last Updated on STN: 20020122  
 Entered Medline: 20011204  
 AB BACKGROUND: CNS-irradiation in prepubertal children with leukemia or brain tumors can lead to precocious or in high doses to delayed puberty. The underlying mechanisms of these disorders are unknown. METHODS: A new animal model of experimentally induced pubertal disorders by cranial irradiation has been developed. In infantile or juvenile (12 - 23 days old) female rats precocious or delayed puberty have been induced by selective cranial Co60-irradiation (4 - 18 Gy). At age of 32 - 38 days or 3 months relevant hormone parameters have been studied basal and after stimulated conditions. RESULTS: Low radiation doses (5 or 6 Gy) led to accelerated onset of puberty as well as elevated LH- and estradiol levels. High radiation doses (9 - 18 Gy) caused retardation of sexual development, lower gonadotropin levels and growth retardation associated with growth hormone deficiency. After cranial irradiation with 5 Gy the release rates of the inhibitory neurotransmitter gamma-aminobutyric-acid (GABA) from hypothalamic explants were significantly lower (p < 0,05). The gonadotropin-releasing-hormone (GnRH) expression in the hypothalamic preoptic area of irradiated animals (5 Gy) was significantly higher than in controls (p < 0,05). CONCLUSION: The GnRH-pulse generator is very radiosensitive as low dose irradiation causes precocious puberty, whereas high dose irradiation is associated with delayed sexual maturation. Radiation induced precocious puberty might be caused by damage to inhibitory GABAergic neurons leading to disinhibition and premature

activation of GnRH neurons. Our animal model of cranial irradiation seems to be suitable to study neurotransmitter disorders, molecular mechanisms and potential preventive intervention of radiation induced pubertal changes.

L11 ANSWER 3 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 1999258039 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 10326189  
TITLE: Effects of puberty on bone age maturation in a girl after medulloblastoma therapy.  
AUTHOR: Marx M; Schoof E; Grabenbauer G G; Beck J D; Doerr H G  
CORPORATE SOURCE: Division of Paediatric Endocrinology, University of Erlangen-Nuremberg, Germany.  
SOURCE: Journal of pediatric and adolescent gynecology, (1999 May) 12 (2) 62-6.  
Journal code: 9610774. ISSN: 1083-3188.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: (CASE REPORTS)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199906  
ENTRY DATE: Entered STN: 19990714  
Last Updated on STN: 19990714  
Entered Medline: 19990629

AB BACKGROUND: Craniospinal radiotherapy for malignant brain tumors can result in a variety of neuroendocrine disturbances, among which are the development of growth hormone deficiency and early puberty, which can markedly reduce adult height. METHODS: The authors report the case of a girl who received craniospinal radiotherapy for a medulloblastoma at the age of 3.4 years. At 9.1 years, growth hormone therapy was started, and spontaneous onset of puberty (Tanner stage B2) occurred at age 10.3 years. Interval until menarche was short, at only 0.9 years. RESULTS: Although chronologic age at appearance of Tanner stages was within the normal range, the patient showed a rapid acceleration in skeletal maturation, resulting in adult short stature. CONCLUSION: Bone age seems to be a more precise parameter for biologic maturation in some patients after craniospinal irradiation than is clinical assessment of pubertal stages. Thus, if progression of bone age and decreasing final height predictions are noted, puberty should be stopped with gonadotropin-releasing hormone analogs, even if pubertal development seems to be adequate for chronologic age, because this increases the remaining time for growth hormone treatment.

L11 ANSWER 4 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 1998419995 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 9749566  
TITLE: Changes in peripheral blood levels and pulse frequencies of GnRH in patients with hypopituitarism.  
AUTHOR: Hayashi M; Takanashi N; Yaoi Y  
CORPORATE SOURCE: Department of Obstetrics and Gynecology, Koshigaya Hospital, Dokkyo University School of Medicine, Japan..  
mhayashi@dokkyomed.ac.jp  
SOURCE: American journal of the medical sciences, (1998 Sep) 316 (3) 213-9.  
Journal code: 0370506. ISSN: 0002-9629.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: (CLINICAL TRIAL)  
(CONTROLLED CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals  
ENTRY MONTH: 199810

ENTRY DATE: Entered STN: 19981020  
Last Updated on STN: 19981020  
Entered Medline: 19981007

AB Pituitary dysfunction occasionally results from **brain tumors** or the surgical resection of **brain tumors**. The authors examined two patients with hypogonadotropic secondary amenorrhea, who had undergone surgical removal of **brain tumors**. Changes in immunoreactive **gonadotropin-releasing hormone** (GnRH) secretion are of interest in patients with a gonadotropin and gonadal steroid deficit, because both steroid and pituitary feedback systems are altered by tumors or tumor resection. The authors thus measured GnRH, luteinizing hormone, and follicle-stimulating hormone levels every 15 minutes for 4 hours by radioimmunoassay and investigated qualitative and quantitative changes in the pulsatile patterns of these hormones in two hypogonadotropic hypogonadism patients. They also performed similar multiple measurements of GnRH in two normal cycle women in follicular phase and two postmenopausal women. The concentration of plasma GnRH in two hypopituitarism patients was compared with that in two normal cycle women and two postmenopausal women. The study showed that the peripheral blood level of GnRH was significantly lower in two hypopituitarism patients than in both normal cycle and postmenopausal women, and that the pulsatile frequency was not different among these three groups. These findings suggest that alteration of feedback systems results in a decrease in the blood level of GnRH, and that pulses of GnRH maintain normal fluctuation despite the alteration of the hormonal circumstances in two hypogonadotropic hypogonadism patients.

L11 ANSWER 5 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 97284740 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 9139717  
TITLE: An alternative **gonadotropin-releasing hormone** (GnRH) RNA splicing product found in cultured GnRH neurons and mouse hypothalamus.  
AUTHOR: Zhen S; Dunn I C; Wray S; Liu Y; Chappell P E; Levine J E; Radovick S  
CORPORATE SOURCE: Department of Medicine, Division of Endocrinology, Children's Hospital, Harvard Medical School, Boston, Massachusetts 02115, USA.  
CONTRACT NUMBER: HD30040 (NICHD)  
SOURCE: Journal of biological chemistry, (1997 May 9) 272 (19) 12620-5.  
Journal code: 2985121R. ISSN: 0021-9258.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199706  
ENTRY DATE: Entered STN: 19970630  
Last Updated on STN: 20021015  
Entered Medline: 19970616

AB **Gonadotropin-releasing hormone** (GnRH) is encoded by the proGnRH gene which contains four exons and three introns. In this study, two immortalized GnRH-expressing cell lines (Gn11 and NLT) were characterized. The NLT and Gn11 cells, derived from a same **brain tumor** in a transgenic mouse, display neuronal morphology and neuron-specific markers. However, NLT cells secrete much higher levels of GnRH than Gn11 cells. To delineate the mechanism underlying this difference, reverse transcriptase-polymerase chain reaction and RNase protection assays were performed to examine proGnRH gene expression. While the mature proGnRH mRNA was predominately expressed in NLT cells, Gn11 cells express an abundant short transcript. Sequence analysis revealed that this short transcript contains exons 1, 3, and 4, but not exon 2, which encodes the GnRH decapeptide. RNase

protection assays demonstrated that NLT cells express much higher levels of mature proGnRH mRNA than Gn11 cells. The lower level of GnRH secreting capacity in Gn11 cells is due, in part, to decreased expression of mature proGnRH mRNA. When proGnRH gene expression in the mouse brain was examined, the same short splicing variant was observed in the olfactory area and preoptic area-anterior hypothalamus. But the prevalent transcript in these regions was the mature proGnRH mRNA. In contrast, only the mature proGnRH mRNA was found in the caudal hypothalamus. These results suggest that alternative splicing may be one of the mechanisms regulating proGnRH gene expression in the animal brain.

L11 ANSWER 6 OF 7 MEDLINE on STN  
 ACCESSION NUMBER: 97395596 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 9251734  
 TITLE: Increased LH and FSH secretion after cranial irradiation in boys.  
 AUTHOR: Lannering B; Jansson C; Rosberg S; Albertsson-Wikland K  
 CORPORATE SOURCE: Department of Paediatrics, University of Goteborg, Sweden.  
 SOURCE: Medical and pediatric oncology, (1997 Oct) 29 (4) 280-7.  
 Journal code: 7506654. ISSN: 0098-1532.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 199709  
 ENTRY DATE: Entered STN: 19970916  
 Last Updated on STN: 19970916  
 Entered Medline: 19970903

AB The effect of high-dose cranial- and craniospinal irradiation and chemotherapy on the gonadotropin-sex steroid axis was studied during different stages of puberty by measuring pulsatile secretion of luteinizing hormone (LH), follicle-stimulating hormone (FSH) and testosterone. The patients were thirteen boys who had been treated for malignant brain tumor residing well away from the hypothalamo-pituitary region. The median time to follow-up was 9 (1-16) years. The onset of puberty was early in the patients, median 10.5 years, compared to the average age for Swedish boys, which is at median 12.4 years. There was, before puberty, no significant difference in LH and FSH secretion between patients and a control group of normal boys. In early, mid- and late stages of puberty, however, LH and FSH secretion was increased in the patients overall, whereas testosterone secretion was maintained within the normal range in spite of signs of gonadotoxicity with small testicular volumes. These results indicate that the vulnerable parts of the gonadotropin releasing hormone (GnRH)-gonadotropin (LH, FSH)-gonadal axis are the regulatory system that determines the timing of pubertal induction and the gonads. The GnRH-LH, FSH-releasing neurons appear relatively resistant to cranial irradiation as they are able to respond with supranormal LH and FSH levels for long periods of time after treatment.

L11 ANSWER 7 OF 7 MEDLINE on STN  
 ACCESSION NUMBER: 95017023 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 7931595  
 TITLE: Precocious puberty in a girl with an hCG-secreting suprasellar immature teratoma. Case report.  
 AUTHOR: Kitanaka C; Matsutani M; Sora S; Kitanaka S; Tanae A; Hibi I  
 CORPORATE SOURCE: Department of Neurosurgery, Tokyo University School of Medicine, Japan.  
 SOURCE: Journal of neurosurgery, (1994 Oct) 81 (4) 601-4.  
 Journal code: 0253357. ISSN: 0022-3085.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: (CASE REPORTS)  
 Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English  
FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals  
ENTRY MONTH: 199410  
ENTRY DATE: Entered STN: 19941222  
Last Updated on STN: 19941222  
Entered Medline: 19941027

AB Although precocious puberty is common in boys with human chorionic gonadotropin (hCG)-secreting brain tumors, it is extremely rare in girls. The authors describe a 6-year-old girl with an hCG-secreting suprasellar immature teratoma who presented with diabetes insipidus, increased intracranial pressure, and precocious puberty. On admission, breast budding was observed. The serum hCG level was 1230 mIU/ml. Both luteinizing hormone (LH) and follicle-stimulating hormone (FSH) remained below detectable levels, even after gonadotropin-releasing hormone stimulation. Serum estrogen and androgen were moderately elevated. After chemotherapy, breast budding disappeared with normalization of serum hCG. It has been believed that hCG does not produce precocious puberty in girls in the absence of FSH, and this has been used as an explanation for the rarity of precocious puberty in girls with hCG-secreting brain tumors. However, it has also been reported that hCG has not only LH activity but also intrinsic, although weak, FSH-like activity. In the present case, this FSH-like activity was considered to have played a role in the development of precocious puberty. It is speculated that a very high level of serum hCG can produce precocious puberty in girls. The rarity of intracranial germ-cell tumors with a high potential of hCG secretion may be one of the reasons why hCG-induced precocious puberty is uncommon in girls.